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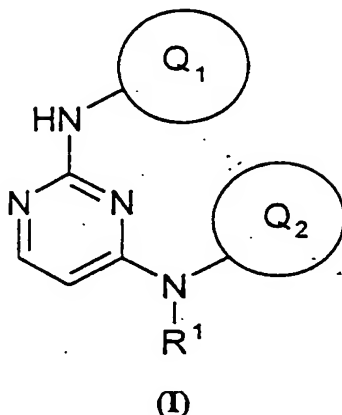
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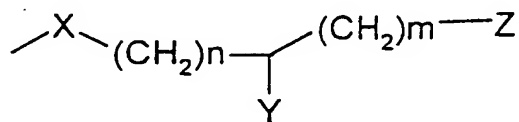
1. A pyrimidine derivative of the formula (I)



wherein

- R<sup>1</sup> is selected from hydrogen, (1-6C)alkyl [optionally substituted by one or two substituents independently selected from halo, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, hydroxy, cyano, (1-4C)alkoxy, (1-4C)alkoxycarbonyl, carbamoyl, -NHCO(1-4C)alkyl, trifluoromethyl, phenylthio, phenoxy, pyridyl, morpholino], benzyl, 2-phenylethyl, (3-5C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent, or one phenyl substituent], N-phthalimido-(1-4C)alkyl, (3-5C)alkynyl [optionally substituted by one phenyl substituent] and (3-6C)cycloalkyl-(1-6C)alkyl;
- wherein any phenyl or benzyl group in R<sup>1</sup> is optionally substituted by up to three substituents independently selected from halogeno, hydroxy, nitro, amino, (1-3C)alkylamino, di-[(1-3C)alkyl]amino, cyano, trifluoromethyl, (1-3C)alkyl [optionally substituted by 1 or 2 substituents independently selected from halogeno, cyano, amino, (1-3C)alkylamino, di-[(1-3C)alkyl]amino, hydroxy and trifluoromethyl], (3-5C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent], (3-5C)alkynyl, (1-3C)alkoxy, -SH, -S-(1-3C)alkyl, carboxy, (1-3C)alkoxycarbonyl;
- Q<sub>1</sub> and Q<sub>2</sub> are independently selected from phenyl, naphthyl, indanyl and 1,2,3,4-tetrahydronaphthyl;

and one or both of  $Q_1$  and  $Q_2$  bears on any available carbon atom one substituent of the formula (Ia) and  $Q_2$  may optionally bear on any available carbon atom further substituents of the formula (Ia)



(Ia)

[provided that when present in  $Q_1$  the substituent of formula (Ia) is not adjacent to the -NH-link];

wherein

X is  $\text{CH}_3$ , O, S, NH or  $\text{NR}_x$  [wherein  $\text{R}_x$  is (1-4C)alkyl, optionally substituted by one

substituent selected from halo, amino, cyano, (1-4C)alkoxy or hydroxy];

Y is H or as defined for Z;

Z is OH, SH,  $\text{NH}_2$ , (1-4C)alkoxy, (1-4C)alkylthio, -NH(1-4C)alkyl, -N[(1-4C)alkyl]<sub>2</sub>, -NH-(3-8C)cycloalkyl, pyrrolidin-1-yl, piperidin-1-yl, piperazin-1-yl [optionally substituted in the 4-position by (1-4C)alkyl or (1-4C)alkanoyl], morpholino or thiomorpholino;

n is 1, 2 or 3; m is 1, 2 or 3;

and  $Q_1$  may optionally bear on any available carbon atom up to four substituents independently selected from halogeno, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],

(2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl, fluoro-(1-4C)alkyl, amino-(1-3C)alkyl, (1-4C)alkylamino-(1-3C)alkyl, di-[(1-4C)alkyl]amino-(1-3C)alkyl, cyano-(1-4C)alkyl, (2-4C)alkanoyloxy-(1-4C)-alkyl, (1-4C)alkoxy-(1-3C)alkyl, carboxy-(1-4C)alkyl, (1-4C)alkoxycarbonyl-(1-4C)alkyl, carbamoyl-(1-4C)alkyl, N-(1-4C)alkylcarbamoyl-(1-4C)alkyl,

N,N-di-[(1-4C)alkyl]-carbamoyl-(1-4C)alkyl, pyrrolidin-1-yl-(1-3C)alkyl, piperidin-1-yl-(1-3C)alkyl, piperazin-1-yl-(1-3C)alkyl, morpholino-(1-3C)alkyl, thiomorpholino-(1-3C)alkyl, piperazin-1-yl, morpholino, thiomorpholino, (1-4C)alkylthio, (1-4C)alkylsulphinyl, (1-4C)alkylsulphonyl, hydroxy-(2-4C)alkylthio, hydroxy-(2-4C)alkylsulphinyl, hydroxy-(2-4C)alkylsulphonyl, ureido ( $\text{H}_2\text{N}-\text{CO}-\text{NH}-$ ),

(1-4C)alkylNH-CO-NH-, di-[(1-4C)alkyl]N-CO-NH-, (1-4C)alkylNH-CO-N[(1-4C)alkyl]-, di-[(1-4C)alkyl]N-CO-N[(1-4C)alkyl]-, carbamoyl, N-[(1-4C)alkyl]carbamoyl, N,N-di-[(1-4C)alkyl]carbamoyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino;

- 5 and also independently, or in addition to, the above substituents, Q<sub>1</sub> may optionally bear on any available carbon atom up to two further substituents independently selected from (3-8C)cycloalkyl, phenyl-(1-4C)alkyl, phenyl-(1-4C)alkoxy, phenylthio, phenyl, naphthyl, benzoyl, benzimidazol-2-yl and a 5- or 6-membered aromatic heterocycle (linked via a ring carbon atom and containing one to three heteroatoms independently selected from oxygen, sulphur and nitrogen); wherein said naphthyl, phenyl, benzoyl, 5- or 6-membered aromatic heterocyclic substituents and the phenyl group in said phenyl-(1-4C)alkyl, phenylthio and phenyl-(1-4C)alkoxy substituents may optionally bear up to five substituents independently selected from halogeno, (1-4C)alkyl and (1-4C)alkoxy;
- 15 and Q<sub>2</sub> may optionally bear on any available carbon atom up to four substituents independently selected from halogeno, hydroxy, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent], (2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl, fluoro-(1-4C)alkyl, amino-(1-3C)alkyl, (1-4C)alkylamino-(1-3C)alkyl, di-[(1-4C)alkyl]amino-(1-3C)alkyl, cyano-(1-4C)alkyl, (2-4C)alkanoyloxy-(1-4C)alkyl, (1-4C)alkoxy-(1-3C)alkyl, carboxy-(1-4C)alkyl, (1-4C)alkoxycarbonyl-(1-4C)alkyl, carbamoyl-(1-4C)alkyl, N-(1-4C)alkylcarbamoyl-(1-4C)alkyl, N,N-di-[(1-4C)alkyl]-carbamoyl-(1-4C)alkyl, pyrrolidin-1-yl-(1-3C)alkyl, piperidin-1-yl-(1-3C)alkyl, piperazin-1-yl-(1-3C)alkyl, morpholino-(1-3C)alkyl, thiomorpholino-(1-3C)alkyl, piperazin-1-yl, morpholino, thiomorpholino, (1-4C)alkoxy, cyano-(1-4C)alkoxy, carbamoyl-(1-4C)alkoxy, N-(1-4C)alkylcarbamoyl-(1-4C)alkoxy, N,N-di-[(1-4C)alkyl]-carbamoyl-(1-4C)alkoxy, 2-aminoethoxy, 2-(1-4C)alkylaminoethoxy, 2-di-[(1-4C)alkyl]aminoethoxy, (1-4C)alkoxycarbonyl-(1-4C)alkoxy, halogeno-(1-4C)alkoxy, 2-hydroxyethoxy, (2-4C)alkanoyloxy-(2-4C)alkoxy, 2-(1-4C)alkoxyethoxy, carboxy-(1-4C)alkoxy,
- 30

(3-5C)alkenyloxy, (3-5C)alkynyloxy, (1-4C)alkylthio, (1-4C)alkylsulphinyl, (1-4C)alkylsulphonyl, hydroxy-(2-4C)alkylthio, hydroxy-(2-4C)alkylsulphinyl, hydroxy-(2-4C)alkylsulphonyl, ureido ( $H_2N-CO-NH-$ ), (1-4C)alkylNH-CO-NH-, di-[(1-4C)alkyl]N-CO-NH-, (1-4C)alkylNH-CO-N[(1-4C)alkyl]-, di-[(1-4C)alkyl]N-CO-N[(1-4C)alkyl]-, carbamoyl, N-[(1-4C)alkyl]carbamoyl, N,N-di-[(1-4C)alkyl]carbamoyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino, and also independently, or in addition to, the above optional substituents,  $Q_2$  may optionally bear on any available carbon atom up to two further substituents independently selected from (3-8C)cycloalkyl, phenyl-(1-4C)alkyl, phenyl-(1-4C)alkoxy, phenylthio, phenyl, naphthyl, benzoyl, phenoxy, benzimidazol-2-yl and a 5- or 6-membered aromatic heterocycle (linked via a ring carbon atom and containing one to three heteroatoms independently selected from oxygen, sulphur and nitrogen); wherein said naphthyl, phenyl, benzoyl, 5- or 6-membered aromatic heterocyclic substituents and the phenyl group in said phenyl-(1-4C)alkyl, phenylthio, phenoxy and phenyl-(1-4C)alkoxy substituents may optionally bear up to five substituents independently selected from halogeno, (1-4C)alkyl and (1-4C)alkoxy; or a pharmaceutically-acceptable salt or in-vivo-hydrolysable ester thereof.

2. A pyrimidine derivative of the formula (I) as claimed in claim 1, wherein  $R^1$  is hydrogen, benzyl, (3-5C)alkynyl, (3-6C)cycloalkyl-(1-6C)alkyl, (1-4C)alkyl [optionally substituted by one or two substituents independently selected from hydroxy, amino, halo, trifluoromethyl and cyano] or (3-5C)alkenyl substituted by one to three halo groups or one phenyl substituent;
- $Q_1$  and  $Q_2$  are independently selected from phenyl, naphthyl, indanyl and 1,2,3,4-tetrahydronaphthyl;
- and one or both of  $Q_1$  and  $Q_2$  bears on any available carbon atom one substituent of the formula (Ia) and  $Q_2$  may optionally bear on any available carbon atom further substituents of the formula (Ia) [provided that when present in  $Q_1$  the substituent of formula (Ia) is not adjacent to the -NH- link];
- X is  $CH_2$ , O, S, NH or  $NR_x$  [wherein  $R_x$  is (1-4C)alkyl, optionally substituted by one substituent selected from halo, amino, cyano, (1-4C)alkoxy or hydroxy];
- Y is H or as defined for Z;

Z is OH, SH, NH<sub>2</sub>, (1-4C)alkoxy, (1-4C)alkylthio, -NH(1-4C)alkyl, -N[(1-4C)alkyl]<sub>2</sub>, -NH-(3-8C)cycloalkyl, pyrrolidin-1-yl, piperidin-1-yl, piperazin-1-yl [optionally substituted in the 4-position by (1-4C)alkyl or (1-4C)alkanoyl], morpholino or thiomorpholino; n is 1, 2 or 3; m is 1, 2 or 3;

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and Q<sub>1</sub> may optionally bear on any available carbon atom up to four substituents independently selected from halogeno, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],

(2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl,

10 fluoro-(1-4C)alkyl, amino-(1-3C)alkyl, (1-4C)alkylamino-(1-3C)alkyl, di-[(1-4C)alkyl]amino-(1-3C)alkyl, cyano-(1-4C)alkyl, (2-4C)alkanoyloxy-(1-4C)-alkyl, (1-4C)alkoxy-(1-3C)alkyl, carboxy-(1-4C)alkyl, (1-4C)alkoxycarbonyl-(1-4C)alkyl, carbamoyl-(1-4C)alkyl, N-(1-4C)alkylcarbamoyl-(1-4C)alkyl,

N,N-di-[(1-4C)alkyl]-carbamoyl-(1-4C)alkyl, pyrrolidin-1-yl-(1-3C)alkyl, piperidin-1-yl-

15 (1-3C)alkyl, piperazin-1-yl-(1-3C)alkyl, morpholino-(1-3C)alkyl, thiomorpholino-(1-3C)alkyl, piperazin-1-yl, morpholino, thiomorpholino, (1-4C)alkylthio, (1-4C)alkylsulphinyl, (1-4C)alkylsulphonyl, hydroxy-(2-4C)alkylthio, hydroxy-(2-4C)alkylsulphinyl, hydroxy-(2-4C)alkylsulphonyl, ureido (H<sub>2</sub>N-CO-NH-), (1-4C)alkylNH-CO-NH-, di-[(1-4C)alkyl]N-CO-NH-, (1-4C)alkylNH-CO-N[(1-4C)alkyl]-, 20 di-[(1-4C)alkyl]N-CO-N[(1-4C)alkyl]-, carbamoyl, N-[(1-4C)alkyl]carbamoyl, N,N-di-[(1-4C)alkyl]carbamoyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino;

and also independently, or in addition to, the above substituents, Q<sub>1</sub> may optionally bear on any available carbon atom up to two further substituents independently selected from

25 (3-8C)cycloalkyl, phenyl-(1-4C)alkyl, phenyl-(1-4C)alkoxy, phenylthio, phenyl, naphthyl, benzoyl, benzimidazol-2-yl and a 5- or 6-membered aromatic heterocycle (linked via a ring carbon atom and containing one to three heteroatoms independently selected from oxygen, sulphur and nitrogen); wherein said naphthyl, phenyl, benzoyl, 5- or 6-membered aromatic heterocyclic substituents and the phenyl group in said phenyl-(1-4C)alkyl, phenylthio and 30 phenyl-(1-4C)alkoxy substituents may optionally bear up to five substituents independently selected from halogeno, (1-4C)alkyl and (1-4C)alkoxy;

and Q<sub>2</sub> may optionally bear on any available carbon atom up to four substituents

independently selected from halogeno, hydroxy, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent], (2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl,

- 5 fluoro-(1-4C)alkyl, amino-(1-3C)alkyl, (1-4C)alkylamino-(1-3C)alkyl, di-[(1-4C)alkyl]amino-(1-3C)alkyl, cyano-(1-4C)alkyl, (2-4C)alkanoyloxy-(1-4C)-alkyl, (1-4C)alkoxy-(1-3C)alkyl, carboxy-(1-4C)alkyl, (1-4C)alkoxycarbonyl-(1-4C)alkyl, carbamoyl-(1-4C)alkyl, N-(1-4C)alkylcarbamoyl-(1-4C)alkyl, N,N-di-[(1-4C)alkyl]-carbamoyl-(1-4C)alkyl, pyrrolidin-1-yl-(1-3C)alkyl, piperidin-1-yl-  
10 (1-3C)alkyl, piperazin-1-yl-(1-3C)alkyl, morpholino-(1-3C)alkyl, thiomorpholino-(1-3C)alkyl, piperazin-1-yl, morpholino, thiomorpholino, (1-4C)alkoxy, cyano-(1-4C)alkoxy, carbamoyl-(1-4C)alkoxy, N-(1-4C)alkylcarbamoyl-(1-4C)alkoxy, N,N-di-[(1-4C)alkyl]-carbamoyl-(1-4C)alkoxy, 2-aminoethoxy, 2-(1-4C)alkylaminoethoxy, 2-di-[(1-4C)alkyl]aminoethoxy, (1-4C)alkoxycarbonyl-(1-4C)alkoxy,  
15 halogeno-(1-4C)alkoxy, 2-hydroxyethoxy, (2-4C)alkanoyloxy-(2-4C)alkoxy, 2-(1-4C)alkoxyethoxy, carboxy-(1-4C)alkoxy, (3-5C)alkenyloxy, (3-5C)alkynyloxy, (1-4C)alkylthio, (1-4C)alkylsulphinyl, (1-4C)alkylsulphonyl, hydroxy-(2-4C)alkylthio, hydroxy-(2-4C)alkylsulphinyl, hydroxy-(2-4C)alkylsulphonyl, ureido (H<sub>2</sub>N-CO-NH-), (1-4C)alkylNH-CO-NH-, di-[(1-4C)alkyl]N-CO-NH-, (1-4C)alkylNH-CO-N[(1-4C)alkyl]-,  
20 di-[(1-4C)alkyl]N-CO-N[(1-4C)alkyl]-, carbamoyl, N-[(1-4C)alkyl]carbamoyl, N,N-di-[(1-4C)alkyl]carbamoyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino,

and also independently, or in addition to, the above optional substituents, Q<sub>2</sub> may optionally bear on any available carbon atom up to two further substituents independently selected from

- 25 (3-8C)cycloalkyl, phenyl-(1-4C)alkyl, phenyl-(1-4C)alkoxy, phenylthio, phenyl, naphthyl, benzoyl, phenoxy, benzimidazol-2-yl and a 5- or 6-membered aromatic heterocycle (linked via a ring carbon atom and containing one to three heteroatoms independently selected from oxygen, sulphur and nitrogen); wherein said naphthyl, phenyl, benzoyl, 5- or 6-membered aromatic heterocyclic substituents and the phenyl group in said phenyl-(1-4C)alkyl,  
30 phenylthio, phenoxy and phenyl-(1-4C)alkoxy substituents may optionally bear up to five substituents independently selected from halogeno, (1-4C)alkyl and (1-4C)alkoxy; or a

pharmaceutically-acceptable salt or in-vivo-hydrolysable ester thereof.

3. A pyrimidine derivative of the formula (I) as claimed in claim 1 or 2, wherein  
R<sup>1</sup> is hydrogen, benzyl, (3-5C)alkynyl, (3-6C)cycloalkyl-(1-6C)alkyl, (1-4C)alkyl [optionally  
5 substituted by one or two substituents independently selected from hydroxy, amino, halo,  
trifluoromethyl and cyano] or (3-5C)alkenyl substituted by one to three halo groups or one  
phenyl substituent;

Q<sub>1</sub> and Q<sub>2</sub> are independently selected from phenyl or indanyl;

and one or both of Q<sub>1</sub> and Q<sub>2</sub> bears on any available carbon atom one substituent of the  
10 formula (Ia) and Q<sub>2</sub> may optionally bear on any available carbon atom further substituents of  
the formula (Ia) [provided that when present in Q<sub>1</sub> the substituent of formula (Ia) is not  
adjacent to the -NH- link];

X is CH<sub>2</sub>, O, S, NH or NR<sub>x</sub> [wherein R<sub>x</sub> is (1-4C)alkyl, optionally substituted by one  
substituent selected from halo, amino, cyano, (1-4C)alkoxy or hydroxy];

15 Y is H or as defined for Z;

Z is OH, SH, NH<sub>2</sub>, (1-4C)alkoxy, (1-4C)alkylthio, -NH(1-4C)alkyl, -N[(1-4C)alkyl]<sub>2</sub>,  
-NH-(3-8C)cycloalkyl, pyrrolidin-1-yl, piperidin-1-yl, piperazin-1-yl [optionally substituted  
in the 4-position by (1-4C)alkyl or (1-4C)alkanoyl], morpholino or thiomorpholino;

n is 1, 2 or 3; m is 1, 2 or 3;

20 and Q<sub>1</sub> may optionally bear on any available carbon atom up to four substituents  
independently selected from halogeno, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally  
substituted by up to three halo substituents, or by one trifluoromethyl substituent],

(2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl,  
fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino;

25 and Q<sub>2</sub> may optionally bear on any available carbon atom up to four substituents  
independently selected from halogeno, hydroxy, thio, nitro, carboxy, cyano, (2-4C)alkenyl  
[optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],  
(2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl,  
fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino,  
30 and also independently, or in addition to, the above optional substituents, Q<sub>2</sub> may optionally  
bear on any available carbon atom up to two further substituents independently selected from



phenylthio, phenyl, phenoxy and benzimidazol-2-yl; or a pharmaceutically-acceptable salt or in-vivo-hydrolysable ester thereof.

4. A pyrimidine derivative of the formula (I) as claimed in any of claims 1 to 3, wherein  
5 R<sup>1</sup> is hydrogen, benzyl, (3-5C)alkynyl, (3-6C)cycloalkyl-(1-6C)alkyl, (1-4C)alkyl [optionally substituted by one or two substituents independently selected from hydroxy, amino, halo, trifluoromethyl and cyano] or (3-5C)alkenyl substituted by one to three halo groups or one phenyl substituent;

Q<sub>1</sub> and Q<sub>2</sub> are independently selected from phenyl or indan-5-yl;

10 and one or both of Q<sub>1</sub> and Q<sub>2</sub> bears on any available carbon atom one substituent of the formula (Ia) and Q<sub>2</sub> may optionally bear on any available carbon atom further substituents of the formula (Ia) [provided that when present in Q<sub>1</sub> the substituent of formula (Ia) is not adjacent to the -NH- link];

X is O; Y is H or OH and Z is -NH(1-4C)alkyl, -N[(1-4C)alkyl]<sub>2</sub>, -NH-(3-8C)cycloalkyl,  
15 pyrrolidin-1-yl or piperazin-1-yl [optionally substituted in the 4-position by (1-4C)alkyl or (1-4C)alkanoyl]; n is 1 or 2 and m is 1 or 2;

and Q<sub>1</sub> may optionally bear on any available carbon atom up to four substituents independently selected from halogeno, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],

20 (2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl, fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino; and Q<sub>2</sub> may optionally bear on any available carbon atom up to four substituents

independently selected from halogeno, hydroxy, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],

25 (2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl, fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino, and also independently, or in addition to, the above optional substituents, Q<sub>2</sub> may optionally bear on any available carbon atom up to two further substituents independently selected from phenylthio, phenyl, phenoxy and benzimidazol-2-yl; or a pharmaceutically-acceptable salt or  
30 in-vivo-hydrolysable ester thereof.

5. A pyrimidine derivative of the formula (I) as claimed in any of claims 1 to 4, wherein  $R^1$  is  $-\text{CH}_2\text{CH}=\text{CHBr}$ ,  $-\text{CH}_2\text{CH}_2\text{CH}_2\text{CF}_3$  or  $-\text{CH}_2\text{CH}=\text{CH-phenyl}$ ;

$Q_1$  and  $Q_2$  are independently selected from phenyl or indan-5-yl;

and one or both of  $Q_1$  and  $Q_2$  bears on any available carbon atom one substituent of the

5 formula (Ia) and  $Q_2$  may optionally bear on any available carbon atom further substituents of the formula (Ia) [provided that when present in  $Q_1$  the substituent of formula (Ia) is not adjacent to the  $-\text{NH-}$  link];

X is O; Y is H or OH and Z is  $-\text{NH}(1-4\text{C})\text{alkyl}$ ,  $-\text{N}[(1-4\text{C})\text{alkyl}]_2$ ,  $-\text{NH-(3-8C)cycloalkyl}$ , pyrrolidin-1-yl or piperazin-1-yl [optionally substituted in the 4-position by (1-4C)alkyl or

10 (1-4C)alkanoyl]; n is 1 or 2 and m is 1 or 2;

and  $Q_1$  may optionally bear on any available carbon atom up to four substituents

independently selected from halogeno, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],

(2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl,

15 fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino;

and  $Q_2$  may optionally bear on any available carbon atom up to four substituents

independently selected from halogeno, hydroxy, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent],

(2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl,

20 fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino,

and also independently, or in addition to, the above optional substituents,  $Q_2$  may optionally bear on any available carbon atom up to two further substituents independently selected from phenylthio, phenyl, phenoxy and benzimidazol-2-yl; or a pharmaceutically-acceptable salt or in-vivo-hydrolysable ester thereof.

25

6. A pyrimidine derivative of the formula (I) as claimed in any of claims 1 to 5, wherein  $R^1$  is  $-\text{CH}_2\text{CH}=\text{CHBr}$ ,  $-\text{CH}_2\text{CH}_2\text{CH}_2\text{CF}_3$  or  $-\text{CH}_2\text{CH}=\text{CH-phenyl}$ ;

$Q_1$  and  $Q_2$  are both phenyl;

$Q_1$  bears on any available carbon atom one substituent of the formula (Ia) [provided that the

30 substituent of formula (Ia) is not adjacent to the  $-\text{NH-}$  link];

X is O; Y is H or OH and Z is -NH(1-4C)alkyl, -N[(1-4C)alkyl]<sub>2</sub>, -NH-(3-8C)cycloalkyl, pyrrolidin-1-yl or piperazin-1-yl [optionally substituted in the 4-position by (1-4C)alkyl or (1-4C)alkanoyl]; n is 1 or 2 and m is 1 or 2;

and Q<sub>1</sub> may optionally bear on any available carbon atom up to four substituents

5 independently selected from halogeno, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent], (2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl, fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino; and Q<sub>2</sub> may optionally bear on any available carbon atom up to four substituents

10 independently selected from halogeno, hydroxy, thio, nitro, carboxy, cyano, (2-4C)alkenyl [optionally substituted by up to three halo substituents, or by one trifluoromethyl substituent], (2-4C)alkynyl, (1-5C)alkanoyl, (1-4C)alkoxycarbonyl, (1-6C)alkyl, hydroxy-(1-6C)alkyl, fluoro-(1-4C)alkyl, amino, (1-4C)alkylamino, di-[(1-4C)alkyl]amino, (2-4C)alkanoylamino, and also independently, or in addition to, the above optional substituents, Q<sub>2</sub> may optionally  
15 bear on any available carbon atom up to two further substituents independently selected from phenylthio, phenyl, phenoxy and benzimidazol-2-yl; or a pharmaceutically-acceptable salt or in-vivo-hydrolysable ester thereof.

7. A pyrimidine derivative of the formula (I) as described in claim 5 or 6, other than that  
20 R<sup>1</sup> is H; or a pharmaceutically-acceptable salt or in-vivo-hydrolysable ester thereof.

8. A pyrimidine derivative of the formula (I) as claimed in claim 1, being:

2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2-bromo-4-methylanilino)pyrimidine;

25 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloroanilino)pyrimidine;

2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(3,4-dichloroanilino)pyrimidine;

2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,4-difluoro-

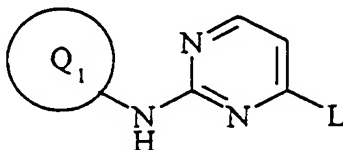
30 (N-cyanomethyl)anilino)pyrimidine;

2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloro-(N-

- 2-fluoroethyl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloro-(*N*-propyn-2-yl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloro-  
 5 (N-cyanomethyl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2-fluoro-5-methylanilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2-cyanoanilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloro-  
 10 (N-2,2-difluoroethyl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloro-(N-4,4,4-trifluorobutyl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2,5-dichloro-(N-3-phenylprop-2-enyl)anilino)pyrimidine;  
 15 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2-fluoro-5-methyl-(N-4,4,4-trifluorobutyl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2-fluoro-5-methyl-(N-3-bromoprop-2-enyl)anilino)pyrimidine;  
 2-{4-[3-(*N,N*-Dimethyl)amino-2-hydroxy-propoxy]anilino}-4-(2-fluoro-5-methyl-(N-3-  
 20 phenylprop-2-enyl)anilino)pyrimidine;  
 or pharmaceutically-acceptable salt or in-vivo hydrolysable ester thereof.

9. A process for the preparation of a compound of the formula (I) as claimed in claim 1, which comprises of a) to h) :-

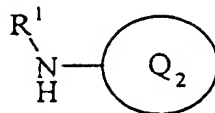
- 25 a) reacting a pyrimidine of formula (II):



(II)

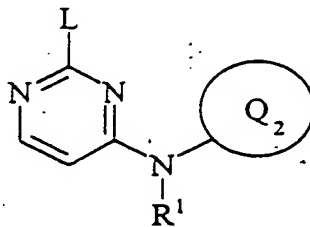
wherein L is a displaceable group as defined below, with a compound of formula (III):

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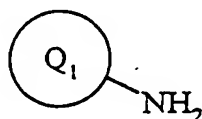
(III)

b) reaction of a pyrimidine of formula (IV):



(IV)

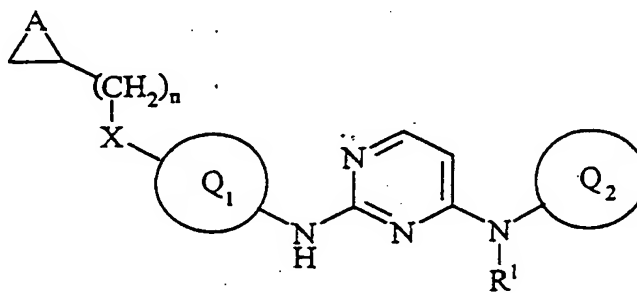
wherein L is a displaceable group as defined below, with a compound of formula (V):



(V)

c) for compounds of formula (I) wherein n is 1, 2 or 3; m = 1 and Y is OH, NH₂ or SH :

reaction of a 3-membered heteroalkyl ring of formula (VI):



(VI)

15 wherein A is O, S or NH;

with a nucleophile of formula (VII):

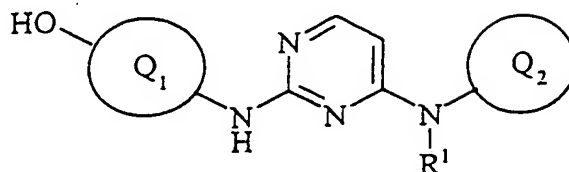
Z-D

(VII)

wherein D is H or a suitable counter-ion;

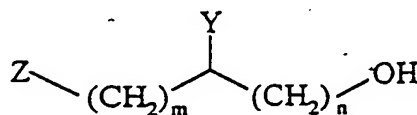
d) for compounds of formula (I) where X is oxygen :

reaction of an alcohol of formula (VIII):



(VIII)

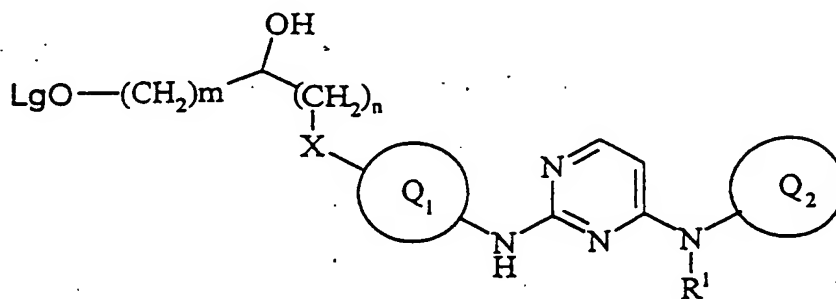
5 with an alcohol of formula (IX):



(IX)

e) for compounds of formula (I) wherein X is CH<sub>2</sub>, O, NH or S; Y is OH and m is 2 or 3 :

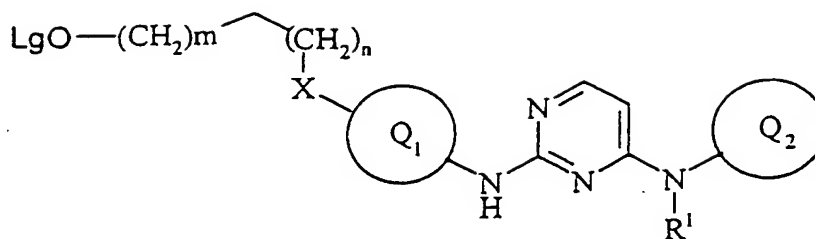
10 reaction of a compound of formula (X) :



(X)

15 wherein -OLg is a leaving group such as mesylate or tosylate; with a nucleophile of formula Z-D (VII) wherein D is H or a suitable counter-ion;

f) for compounds of formula (I) wherein X is CH<sub>2</sub>, O, NH or S; Y is H; n is 1, 2 or 3 and m is 1, 2 or 3 : reaction of a compound of formula (XI) :

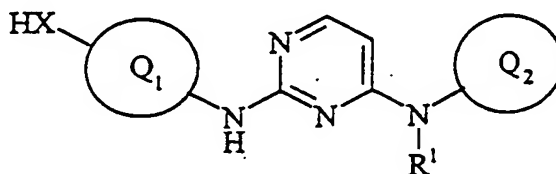


(XI)

wherein -OLg is a leaving group such as mesylate or tosylate; with a nucleophile of formula

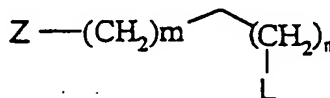
5 Z-D (VII) wherein D is H or a suitable counter-ion;

g) for compounds of formula (I) wherein X is O, NH or S; Y is H; n is 1, 2 or 3 and m is 1, 2 or 3 : reaction of a compound of formula (XII) with a compound of formula (XIII) :



(XII)

10



(XIII)

or

15 h) for compounds of formula (I) in which Z is SH, by conversion of a thioacetate group in a corresponding compound; and thereafter if necessary:

i) converting a compound of the formula (I) into another compound of the formula (I);

ii) removing any protecting groups;

iii) forming a pharmaceutically acceptable salt or *in vivo* hydrolysable ester; wherein L is a

20 displaceable group and D is hydrogen or a counter-ion.

10. A method for producing an anti-cancer effect in a warm blooded animal which comprises administering to said animal an effective amount of a compound of the formula (I)

as claimed in claims 1 to 8, or a pharmaceutically-acceptable salt, or in-vivo hydrolysable ester thereof.

11. A compound of the formula (I) as claimed in claims 1 to 8, or a pharmaceutically-  
5 acceptable salt, or in-vivo hydrolysable ester thereof, for use as a medicament.

12. The use of a compound of the formula (I) as claimed in claims 1 to 8, or a  
pharmaceutically-acceptable salt, or in-vivo hydrolysable ester thereof, in the manufacture of  
a medicament for use in the production of an anti-cancer effect in a warm blooded animal.

10

13. A pharmaceutical composition which comprises a compound of the formula (I) as  
claimed in claims 1 to 8, or a pharmaceutically-acceptable salt or an in-vivo hydrolysable  
ester thereof, and a pharmaceutically-acceptable diluent or carrier.